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## CLAIMS

We claim:

- 1. An isolated caspase or procaspase expressed in immature thymocytes, or an active derivative or fragment thereof wherein said caspase is necessary for apoptosis.
- An isolated caspase according to Claim 1, wherein said caspase is characterized by its ability to be triggered by TCR stimulation with peptide/MHC, anti-CD3ε or other anti-TCR-specific monoclonal antibody, or corticosteroids in thymocytes.
  - 3. The isolated caspase or procaspase of Claim 1, wherein said caspase is a fragment possessing caspase activity.
- 15 4. The isolated caspase or procaspase of Claim 1, wherein said caspase or procaspase is a derivative possessing substantial sequence identity with the endogenous caspase or procaspase.
- 5. The isolated caspase or prodaspase of Claim 1, wherein said caspase or procaspase has the same amino acid sequence as the endogenous caspase or procaspase.
  - 6. The isolated caspase or procaspase of Claim 5, wherein said caspase or procaspase is isolated from immature thymocytes.
- 25 7. The isolated caspase or procaspase of Claim 6, wherein said caspase or procaspase is purified to homogeneity.

- 8. The isolated caspase or procaspase of Claim 6, wherein said caspase or procaspase is substantially free of other thymocyte proteins.
- 9. An isolated nucleic acid molecule which encodes a

  5 caspase or procaspase expressed in immature thymocytes
  and necessary for apoptosis, or an active derivative
  or fragment thereof.
- 10. An isolated nucleic acid molecule according to Claim
   9, wherein the caspase is characterized by its ability
   10 to be triggered by TCR stimulation with peptide/MHC,
   anti-CD3ε or other anti-TCR-specific monoclonal
   antibody, or corticosteroids in thymocytes.
  - 11. The isolated nucleic acid molecule of Claim 9, wherein said molecule comprises at least about 25 nucleotides.
- 15 12. The isolated nucleic acid molecule of Claim 9, wherein said molecule comprises at least about 50 nucleotides.
  - 13. The isolated nucleic acid molecule of Claim 9, wherein said molecule comprises at least about 200 nucleotides.
- 20 14. The isolated nucleic acid molecule of Claim 9, wherein the caspase or procaspase is a derivative possessing substantial sequence identity with the endogenous caspase or procaspase.
- 15. The isolated nucleic acid molecule Claim 9, wherein said caspase or procaspase has the same amino acid sequence as the endogenous caspase or procaspase.



- 16. The isolated nucleic acid molecule Claim 15, wherein said nucleic acid molecule has the same nucleotide sequence as the endogenous gene encoding the caspase or procaspase.
- 5 17. A DNA construct comprising the isolated nucleic acid molecule of Claim 9 operatively linked to a regulatory sequence.
  - 18. A DNA construct comprising the isolated nucleic acid molecule of Claim 15 operatively linked to a regulatory sequence.
  - 19. A DNA construct comprising the isolated nucleic acid molecule of Claim 16 operatively linked to a regulatory sequence.
- 20. A recombinant host cell comprising the isolated

  15 nucleic acid molecule of Claim 9 operatively linked to
  a regulatory sequence.
  - 21. A recombinant host cell comprising the isolated nucleic acid molecule of Claim 15 operatively linked to a regulatory sequence.
- 20 22. A recombinant host cell comprising the isolated nucleic acid molecule of Claim 16 operatively linked to a regulatory sequence.
- 23. The recombinant host cell of Claim 22 wherein said cell is selected from the group consisting of bacterial cells, fungal cells, plant cells, insect cells and mammalian cells.

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- 24. A method for preparing a caspase or procaspase expressed in immature thymocytes, or an active derivative or fragment thereof, wherein said caspase is necessary for apoptosis, comprising culturing the recombinant host cell of Claim 20.
- 25. A method according to Claim 24, wherein the caspase is characterized by its ability to be triggered by TCR stimulation with peptide/MHC, anti-CD3 c or other anti-TCR-specific monoclonal antibody, or corticosteroids in thymocytes
- 26. An antibody, or an antigen-binding fragment thereof, which selectively binds to the caspase or procaspase according to Claim 1 or an active derivative or fragment thereof.
- 15 27. The antibody according to Claim 26, wherein said antibody is a monoclonal antibody.
  - 28. A method for assaying the presence of a caspase or procaspase according to Claim 1 in a cell, comprising contacting said cell with an antibody which specifically binds to the caspase or procaspase.
  - 29. The method of Claim 28, wherein said cell is in a tissue sample.
  - 30. A method of identifying an agent which inhibits the caspase according to Claim 1, comprising the steps of:
- 25 (a) contacting the caspase, or an active derivative or fragment thereof, with a



caspase substrate in the presence of
the agent; and

- (b) identifying inhibition of caspase activity.
- 31. An agent which inhibits caspase activity identified according to the method of Claim 30.
  - 32. A method of identifying an agent which inhibits the caspase according to Claim 1, comprising the steps of:
    - (a) contacting a thymocyte or a cell lysate thereof comprising the caspase or procaspase with the agent; and
    - (b) identifying inhibition of caspase activity.
  - 33. An agent which inhibits caspase activity identified according to the method of Claim 32.
- 34. A method of inhibiting the paspase according to Claim
  15 1, comprising contacting said caspase with an agent
  that inhibits the activity of the caspase.
  - 35. A method of inhibiting apoptosis in a lymphocyte comprising contacting said lymphocyte with an agent which inhibits the caspase according to Claim 1.
- 20 36. A method according to Claim 35, wherein said apoptosis is induced by a member of the group consisting of an antigen, a corticosteroid and anti-CD3 monoclonal antibody.
- 37. A method according to Claim 35, wherein the lymphocyte is an immature thymocyte.

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- 38. A method according to Claim 35, wherein the agent is a tripeptide or tetrapeptide having an amino acid sequence selected from the group consisting of VAD, YVAD, and DEVD, and wherein the tetrapeptide or tripeptide is linked to a compound selected from the group consisiting of fluoromethylketone, acyloxymethylketone, chlormethylketone, diazomethylketone, aldehydes, semicarbazones, nitriles and epoxides.
- 10 39. A method of inhibiting apoptosis in an immature thymocyte comprising contacting said thymocyte with an agent according to Claim 31.
  - 40. A method of inhibiting apoptosis in an immature thymocyte comprising contacting said thymocyte with an agent according to Claim 33.
    - 41. A method of identifying an agent which enhances the caspase according to Claim 1, comprising the steps of:
      - (a) contacting the caspase, or an active derivative or fragment thereof, with a caspase substrate in the presence of the agent; and
      - (b) identifying enhancement of caspase activity.
    - 42. An agent which enhances caspase activity identified according to the method of claim 41.
- 25 43. A method of identifying an agent which enhances the caspase according to Claim 1, comprising the steps of:

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- (a) contacting a thymocyte or a cell lysate thereof comprising the caspase or procaspase, with the agent; and
- (b) identifying shancement of caspase activity.
- 5 44. An agent which enhances caspase activity identified according to the method claim 43.
  - 45. A method of enhancing the caspase according to Claim 1, comprising contacting said caspase with an agent that enhances the activity of the caspase.
- 10 46. A method of enhancing apoptosis in a lymphocyte comprising contacting said lymphocyte with an agent which enhances the caspase according to Claim 1.
  - 47. A method according to Claim 46, wherein the lymphocyte is an immature thymocyte.
- 15 48. A method of enhancing apoptosis in an immature thymocyte comprising contacting said thymocyte with an agent according to Claim 42
- 49. A method of treating an auto mmune disease in a mammal comprising administering to the mammal an effective amount of an agent which enhances the activity of the caspase according to Claim 1.
  - 50. The method of Claim 49, wherein the autoimmune disease is chronic hepatitis or diabetes mellitus.

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- 51. A method of enhancing an immune response against an antigen in a mammal comprising administering to the mammal an effective amount of an agent which inhibits the activity of the caspase according to Claim 1 and an antigen.
- 52. A method of treating a cancer in a mammal comprising administering to the mammal an effective amount of an agent which inhibits the activity of the caspase according to Claim 1 and a cancer antigen.

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